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<120> TREATMENT FOR INSULIN DEPENDENT DIABETES

<130> 10274-008003 <140> US 09/234,290 <141> 1999-01-20 <150> US 08/447,118 <151> 1995-05-22 <150> US 08/029,330 <151> 1993-02-09 <150> PCT/US94/01456 <151> 1994-02-09 <160> 18 <170> FastSEQ for Windows Version 4.0 <210> 1 <211> 360 <212> DNA <213> Homo sapiens <220> <221> misc_feature <222> (0)...(0) <223> pBAG159 insert: HP1/2 heavy chain variable region; amino acid 1 is Glu (E) but Gln (Q) may be substituted <221> CDS <222> (1) ... (360) <400> 1 gtc aaa ctg cag cag tct ggg gca gag ctt gtg aag cca ggg gcc tca 48 Val Lys Leu Gln Gln Ser Gly Ala Glu Leu Val Lys Pro Gly Ala Ser gtc aag ttg tcc tgc aca gct tct ggc ttc aac att aaa gac acc tat 96 Val Lys Leu Ser Cys Thr Ala Ser Gly Phe Asn Ile Lys Asp Thr Tyr atg cac tgg gtg aag cag agg cct gaa cag ggc ctg gag tgg att gga 144 Met His Trp Val Lys Gln Arg Pro Glu Gln Gly Leu Glu Trp Ile Gly 35 45 agg att gat cct gcg agt ggc gat act aaa tat gac ccg aag ttc cag 192 Arg Ile Asp Pro Ala Ser Gly Asp Thr Lys Tyr Asp Pro Lys Phe Gln

50 55 60 gto aag goo act att aca gog gao acg too too aac aca goo tgg otg 240 Val Lys Ala Thr Ile Thr Ala Asp Thr Ser Ser Asn Thr Ala Trp Leu 288 cag ctc agc agc ctg aca tct gag gac act gcc gtc tac tac tgt gca Gln Leu Ser Ser Leu Thr Ser Glu Asp Thr Ala Val Tyr Tyr Cys Ala 85 gac gga atg tgg gta tca acg gga tat gct ctg gac ttc tgg ggc caa 336 Asp Gly Met Trp Val Ser Thr Gly Tyr Ala Leu Asp Phe Trp Gly Gln ggg acc acg gtc acc gtc tcc tca 360 Gly Thr Thr Val Thr Val Ser Ser 115 <210> 2 <211> 120 <212> PRT <213> Homo sapiens <400> 2 Val Lys Leu Gln Gln Ser Gly Ala Glu Leu Val Lys Pro Gly Ala Ser 10 Val Lys Leu Ser Cys Thr Ala Ser Gly Phe Asn Ile Lys Asp Thr Tyr 25 Met His Trp Val Lys Gln Arg Pro Glu Gln Gly Leu Glu Trp Ile Gly Arg Ile Asp Pro Ala Ser Gly Asp Thr Lys Tyr Asp Pro Lys Phe Gln Val Lys Ala Thr Ile Thr Ala Asp Thr Ser Ser Asn Thr Ala Trp Leu 70 75 Gln Leu Ser Ser Leu Thr Ser Glu Asp Thr Ala Val Tyr Tyr Cys Ala Asp Gly Met Trp Val Ser Thr Gly Tyr Ala Leu Asp Phe Trp Gly Gln 100 105 Gly Thr Thr Val Thr Val Ser Ser <210> 3 <211> 318 <212> DNA <213> Homo sapiens <220> <221> misc_feature <222> (0)...(0) <223> pBAG172 insert: HP1/2 light chain variable region

<400> 3

<221> CDS

<222> (1)...(318)

<223> HP1/2 light chain variable region

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_		_				_	_	_	_	_	_			aat Asn	_		96
_	_				_	_			_				_	ctg Leu		נ	L44
		_			_			-	_		_	_		act Thr		1	192
_				_	_									cag Gln	_	2	240
_	_	_	_	_			_	_	_	_		_		ccg Pro 95		2	288
	ttc Phe					_	_		_							3	318

210> 4 <211> 106 <212> PRT

<213> Homo sapiens

<400> 4

 Ser
 Ile
 Val
 Met
 Thr
 Gln
 Thr
 Pro
 Lys
 Phe
 Leu
 Leu
 Val
 Ser
 Ala
 Gly

 Asp
 Arg
 Val
 Thr
 Ile
 Thr
 Cys
 Lys
 Ala
 Ser
 Gln
 Ser
 Val
 Thr
 Asn
 Asp
 Asp

 Val
 Ala
 Try
 Tyr
 Gln
 Gln
 Lys
 Pro
 Gln
 Ser
 Pro
 Leu
 Leu
 Ile

 Jan
 Ala
 Ser
 Asn
 Arg
 Tyr
 Thr
 Gly
 Val
 Pro
 Lys
 Leu
 Leu
 Ile
 Ile
 Asn
 Asp
 Asp
 Asp
 Pro
 Tyr
 Asp
 Arg
 Phe
 Thr
 Brown
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 Arg
 Phe
 Thr
 Pro
 Asp
 Arg
 Phe
 Thr
 Phe
 Thr
 Ile
 Ser
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 Thr
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 Ala
 Ala
 Ala
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100 105

<210> 5 <211> 429

<212> DNA

<213> Homo sapiens

<220> <221> CDS

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<222> (1) . . . (57)
<221> mat_peptide
<222> (58)...(429)
<221> misc feature
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                                                                        48
Met Asp Trp Thr Trp Arg Val Phe Cys Leu Leu Ala Val Ala Pro Gly
gee cae tee cag gte caa etg cag gag age ggt eea ggt ett gtg aga
                                                                        96
Ala His Ser Gln Val Gln Leu Gln Glu Ser Gly Pro Gly Leu Val Arg
cet age cag ace etg age etg ace tge ace geg tet gge tte aac att
                                                                       144
Pro Ser Gln Thr Leu Ser Leu Thr Cys Thr Ala Ser Gly Phe Asn Ile
     15
                         20
aaa gac acc tat atg cac tgg gtg aga cag cca cct gga cga ggt ctt
                                                                       192
Lys Asp Thr Tyr Met His Trp Val Arg Gln Pro Pro Gly Arg Gly Leu
30
gag tgg att gga agg att gat cct gcg agt ggc gat act aaa tat gac
                                                                       240
Glu Trp Ile Gly Arg Ile Asp Pro Ala Ser Gly Asp Thr Lys Tyr Asp
                 50
                                                          60
ccg aag ttc cag gtc aga gtg aca atg ctg gta gac acc agc agc aac
                                                                       288
Pro Lys Phe Gln Val Arg Val Thr Met Leu Val Asp Thr Ser Ser Asn
cag ttc agc ctg aga ctc agc agc gtg aca gcc gcc gac acc gcg gtc
                                                                       336
Gln Phe Ser Leu Arg Leu Ser Ser Val Thr Ala Ala Asp Thr Ala Val
                                                                       384
tat tat tgt gca gac gga atg tgg gta tca acg gga tat gct ctg gac
Tyr Tyr Cys Ala Asp Gly Met Trp Val Ser Thr Gly Tyr Ala Leu Asp
tte tgg gge caa ggg ace acg gte ace gte tee tea ggt gag tee
                                                                       429
Phe Trp Gly Gln Gly Thr Thr Val Thr Val Ser Ser Gly Glu Ser
110
                    115
                                         120
<210> 6
<211> 143
<212> PRT
<213> Homo sapiens
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<400> 6
Met Asp Trp Thr Trp Arg Val Phe Cys Leu Leu Ala Val Ala Pro Gly
                -15
                                     -10
Ala His Ser Gln Val Gln Leu Gln Glu Ser Gly Pro Gly Leu Val Arg
Pro Ser Gln Thr Leu Ser Leu Thr Cys Thr Ala Ser Gly Phe Asn Ile
Lys Asp Thr Tyr Met His Trp Val Arg Gln Pro Pro Gly Arg Gly Leu
                    35
                                         40
Glu Trp Ile Gly Arg Ile Asp Pro Ala Ser Gly Asp Thr Lys Tyr Asp
                                     55
                50
Pro Lys Phe Gln Val Arg Val Thr Met Leu Val Asp Thr Ser Ser Asn
                                 70
Gln Phe Ser Leu Arg Leu Ser Ser Val Thr Ala Ala Asp Thr Ala Val
                             85
Tyr Tyr Cys Ala Asp Gly Met Trp Val Ser Thr Gly Tyr Ala Leu Asp
                                             105
Phe Trp Gly Gln Gly Thr Thr Val Thr Val Ser Ser Gly Glu Ser
                    115
<210> 7
<211> 386
<212> DNA
<213> Homo sapiens
<220>
<221> CDS
<222> (1) . . . (384)
<221> sig peptide
<222> (1)...(57)
<221> mat_peptide
<222> (58)...(384)
<221> misc_feature
<222> (0)...(0)
<223> pBAG198 insert: VK2 (SVMDY) light chain variable
      region
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atg ggt tgg tcc tgc atc atc ctg ttc ctg gtt gct acc gct acc ggt
                                                                        48
Met Gly Trp Ser Cys Ile Ile Leu Phe Leu Val Ala Thr Ala Thr Gly
                -15
                                     -10
                                                                        96
gtc cac tcc agc atc gtg atg acc cag agc cca agc agc ctg agc gcc
Val His Ser Ser Ile Val Met Thr Gln Ser Pro Ser Ser Leu Ser Ala
            1
age gtg ggt gac aga gtg ace ate ace tgt aag gee agt cag agt gtg
                                                                       144
Ser Val Gly Asp Arg Val Thr Ile Thr Cys Lys Ala Ser Gln Ser Val
     15
act aat gat gta gct tgg tac cag cag aag cca ggt aag gct cca aag
                                                                       192
Thr Asn Asp Val Ala Trp Tyr Gln Gln Lys Pro Gly Lys Ala Pro Lys
```

```
35
                                          40
                                                               45
 30
                                                                       240
ctg ctg atc tac tat gca tcc aat cgc tac act ggt gtg cca gat aga
Leu Leu Ile Tyr Tyr Ala Ser Asn Arg Tyr Thr Gly Val Pro Asp Arg
                 50
                                      55
                                                                       288
ttc agc ggt agc ggt tat ggt acc gac ttc acc ttc acc atc agc agc
Phe Ser Gly Ser Gly Tyr Gly Thr Asp Phe Thr Phe Thr Ile Ser Ser
ctc cag cca gag gac atc gcc acc tac tac tgc cag cag gat tat agc
                                                                       336
Leu Gln Pro Glu Asp Ile Ala Thr Tyr Tyr Cys Gln Gln Asp Tyr Ser
tot cog tac acg ttc ggc caa ggg acc aag gtg gaa atc aaa cgt aag
                                                                       384
Ser Pro Tyr Thr Phe Gly Gln Gly Thr Lys Val Glu Ile Lys Arg Lys
     95
                                                                       386
tg
<210> 8
<211> 128
<212> PRT
<213> Homo sapiens
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<221> SIGNAL
<222> (1)...(19)
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Met Gly Trp Ser Cys Ile Ile Leu Phe Leu Val Ala Thr Ala Thr Gly
                -15
                                     -10
Val His Ser Ser Ile Val Met Thr Gln Ser Pro Ser Ser Leu Ser Ala
Ser Val Gly Asp Arg Val Thr Ile Thr Cys Lys Ala Ser Gln Ser Val
Thr Asn Asp Val Ala Trp Tyr Gln Gln Lys Pro Gly Lys Ala Pro Lys
Leu Leu Ile Tyr Tyr Ala Ser Asn Arg Tyr Thr Gly Val Pro Asp Arg
Phe Ser Gly Ser Gly Tyr Gly Thr Asp Phe Thr Phe Thr Ile Ser Ser
                                 70
Leu Gln Pro Glu Asp Ile Ala Thr Tyr Tyr Cys Gln Gln Asp Tyr Ser
Ser Pro Tyr Thr Phe Gly Gln Gly Thr Lys Val Glu Ile Lys Arg Lys
    95
                        100
                                             105
<210> 9
<211> 1347
<212> DNA
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<220>
<221> CDS
<222> (1) ... (1338)
<221> misc feature
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<222> (1)...(219)

<223> VCAM-1 gene segment: This portion of the sequence corresponds, in part, to Exons I, II and III nucleotide sequence of Cybulsky et al. Proc. Nat'l. Acad. Sci. USA 88: 7861 (1991).

- <221> misc_feature
- <222> (220)...(229)
- <223> Hinge region: This portion of the sequence corresponds, in part, to Exons I, II and III nucleotide sequence of Cybulsky et al. Proc. Nat'l. Acad. Sci. USA 88: 7861 (1991).
- <221> misc feature
- <222> (230) . . . (338)
- <223> Heavy chain constant region 2: This portion of the sequence corresponds, in part, to Fig. 12A in PCT/US92/02050 and represents the heavy chain constant region 2 of Human IgGl heavy chain constant region.
- <221> misc_feature
- <222> (339)...(446)
- <223> Heavy chain constant region 3: This portion of the sequence corresponds, in part, to Fig. 12A in PCT/US92/02050 and represents the heavy chain constant region 3 of Human IgGl heavy chain constant region.

<400> 9

atg cct ggg aag atg gtc gtg atc ctt gga gcc tca aat ata ctt tgg Met Pro Gly Lys Met Val Val Ile Leu Gly Ala Ser Asn Ile Leu Trp ata atg ttt gca gct tct caa gct ttt aaa atc gag acc acc cca gaa 96 Ile Met Phe Ala Ala Ser Gln Ala Phe Lys Ile Glu Thr Thr Pro Glu 25

48

tet aga tat ett get eag att ggt gae tee gte tea ttg aet tge age 144 Ser Arg Tyr Leu Ala Gln Ile Gly Asp Ser Val Ser Leu Thr Cys Ser

acc aca ggc tgt gag tcc cca ttt ttc tct tgg aga acc cag ata gat 192 Thr Thr Gly Cys Glu Ser Pro Phe Phe Ser Trp Arg Thr Gln Ile Asp

240 agt cca ctg aat ggg aag gtg acg aat gag ggg acc aca tct acg ctg Ser Pro Leu Asn Gly Lys Val Thr Asn Glu Gly Thr Thr Ser Thr Leu 65 70

288 aca atg aat cct gtt agt ttt ggg aac gaa cac tct tac ctg tgc aca Thr Met Asn Pro Val Ser Phe Gly Asn Glu His Ser Tyr Leu Cys Thr 85

336 gca act tgt gaa tct agg aaa ttg gaa aaa gga atc cag gtg gag atc Ala Thr Cys Glu Ser Arg Lys Leu Glu Lys Gly Ile Gln Val Glu Ile

100 105 110 tac tot tit cot aag gat coa gag att cat tig agt ggc cot cig gag 384 Tyr Ser Phe Pro Lys Asp Pro Glu Ile His Leu Ser Gly Pro Leu Glu 120 gct ggg aag ccg atc aca gtc aag tgt tca gtt gct gat gta tac cca 432 Ala Gly Lys Pro Ile Thr Val Lys Cys Ser Val Ala Asp Val Tyr Pro 135 ttt gac agg ctg gag ata gac tta ctg aaa gga gat cat ctc atg aag 480 Phe Asp Arg Leu Glu Ile Asp Leu Leu Lys Gly Asp His Leu Met Lys 150 agt cag gaa ttt ctg gag gat gca gac agg aag tcc ctg gaa acc aag 528 Ser Gln Glu Phe Leu Glu Asp Ala Asp Arg Lys Ser Leu Glu Thr Lys 165 170 agt ttg gaa gta acc ttt act cct gtc att gag gat att gga aaa gtt 576 Ser Leu Glu Val Thr Phe Thr Pro Val Ile Glu Asp Ile Gly Lys Val 180 ctt gtt tgc cga gct aaa tta cac att gat gaa atg gat tct gtg ccc 624 Leu Val Cys Arg Ala Lys Leu His Ile Asp Glu Met Asp Ser Val Pro 200 aca gta agg cag gct gta aaa gaa ttg caa gtc gac aaa act cac aca 672 Thr Val Arg Gln Ala Val Lys Glu Leu Gln Val Asp Lys Thr His Thr tgc cca ccg tgc cca gca cct gaa ctc ctg ggg gga ccg tca gtc ttc 720 Cys Pro Pro Cys Pro Ala Pro Glu Leu Leu Gly Gly Pro Ser Val Phe 225 cto tto coo coa aaa coo aag gao aco oto atg ato too ogg aco oot 768 Leu Phe Pro Pro Lys Pro Lys Asp Thr Leu Met Ile Ser Arg Thr Pro 245 gag gtc aca tgc gtg gtg gtg gac gtg agc cac gaa gac cct gag gtc 816 Glu Val Thr Cys Val Val Val Asp Val Ser His Glu Asp Pro Glu Val 260 aag ttc aac tqq tac qtq qac qtc qtq qaq qtq cat aat qcc aaq aca 864 Lys Phe Asn Trp Tyr Val Asp Gly Val Glu Val His Asn Ala Lys Thr 275 280 aag ceg egg gag gag eag tac aac agc acg tac egg gtg gtc agc gtc 912 Lys Pro Arg Glu Glu Gln Tyr Asn Ser Thr Tyr Arg Val Val Ser Val 295 ctc acc gtc ctg cac cag gac tgg ctg aat ggc aag gag tac aag tgc 960 Leu Thr Val Leu His Gln Asp Trp Leu Asn Gly Lys Glu Tyr Lys Cys 1008 aag gtc tcc aac aaa gcc ctc cca gcc ccc atc gag aaa acc atc tcc Lys Val Ser Asn Lys Ala Leu Pro Ala Pro Ile Glu Lys Thr Ile Ser

El

	_			_	ccc Pro	_	_		_				_			1056
		_		_	acc Thr	_			-							1104
					agc Ser	-										1152
					tac Tyr 390											1200
					tac Tyr											1248
_	_			-	ttc Phe		_			-			_	_		1296
			_	_	aag Lys	_			-		_					1338
tgag	gtgcg	gg														1347
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<400> 10														24		
tcgtcgacaa aactcacaca tgcc 24																
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	<400> 11															
												24				

<210> 12 <211> 115

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<223> oligonucleotide for PCR
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tgctcgagct ctagatatcg attccatgga tcctcacatc ccaatccgcg gccgc
                                                                        115
<210> 13
<211> 41
<212> DNA
<213> Artificial Sequence
<220>
<223> oligonucleotide for PCR
<400> 13
gagctcgagg cggccgcacc atgcctggga agatggtcgt g
                                                                          41
<210> 14
<211> 23
<212> DNA
<213> Artificial Sequence
<220>
<223> oligonucleotide for PCR
<#00> 14
aagtcgactt gcaattcttt tac
                                                                          23
<210> 15
<211> 14
<212> DNA
<213> Artificial Sequence
<220>
<223> oligonucleotide for PCR
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tcgacgcggc cgcg
                                                                         14
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<212> PRT
<213> Homo sapiens
<400> 16
Met Pro Gly Lys Met Val Val Ile Leu Gly Ala Ser Asn Ile Leu Trp
                 5
                                     10
Ile Met Phe Ala Ala Ser Gln Ala Phe Lys Ile Glu Thr Thr Pro Glu
            20
                                 25
                                                      30
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Ser Arg Tyr Leu Ala Gln Ile Gly Asp Ser Val Ser Leu Thr Cys Ser
                            40
Thr Thr Gly Cys Glu Ser Pro Phe Phe Ser Trp Arg Thr Gln Ile Asp
Ser Pro Leu Asn Gly Lys Val Thr Asn Glu Gly Thr Thr Ser Thr Leu
Thr Met Asn Pro Val Ser Phe Gly Asn Glu His Ser Tyr Leu Cys Thr
                                    90
Ala Thr Cys Glu Ser Arg Lys Leu Glu Lys Gly Ile Gln Val Glu Ile
                                105
Tyr Ser Phe Pro Lys Asp Pro Glu Ile His Leu Ser Gly Pro Leu Glu
                            120
Ala Gly Lys Pro Ile Thr Val Lys Cys Ser Val Ala Asp Val Tyr Pro
                        135
                                            140
Phe Asp Arg Leu Glu Ile Asp Leu Leu Lys Gly Asp His Leu Met Lys
Ser Gln Glu Phe Leu Glu Asp Ala Asp Arg Lys Ser Leu Glu Thr Lys
                                    170
Ser Leu Glu Val Thr Phe Thr Pro Val Ile Glu Asp Ile Gly Lys Val
            180
                                185
                                                    190
Leu Val Cys Arg Ala Lys Leu His Ile Asp Glu Met Asp Ser Val Pro
                            200
                                                205
Thr Val Arg Gln Ala Val Lys Glu Leu Gln Val Asp Lys Thr His Thr
                        215
                                            220
Cys Pro Pro Cys Pro Ala Pro Glu Leu Leu Gly Gly Pro Ser Val Phe
                    230
                                        235
Leu Phe Pro Pro Lys Pro Lys Asp Thr Leu Met Ile Ser Arg Thr Pro
                                   250
                245
Glu Val Thr Cys Val Val Val Asp Val Ser His Glu Asp Pro Glu Val
                                265
Lys Phe Asn Trp Tyr Val Asp Gly Val Glu Val His Asn Ala Lys Thr
                            280
Lys Pro Arg Glu Glu Gln Tyr Asn Ser Thr Tyr Arg Val Val Ser Val
                        295
                                            300
Leu Thr Val Leu His Gln Asp Trp Leu Asn Gly Lys Glu Tyr Lys Cys
                    310
                                        315
Lys Val Ser Asn Lys Ala Leu Pro Ala Pro Ile Glu Lys Thr Ile Ser
                325
                                    330
Lys Ala Lys Gly Gln Pro Arg Glu Pro Gln Val Tyr Thr Leu Pro Pro
                                345
Ser Arg Asp Glu Leu Thr Lys Asn Gln Val Ser Leu Thr Cys Leu Val
                            360
                                                365
Lys Gly Phe Tyr Pro Ser Asp Ile Ala Val Glu Trp Glu Ser Asn Gly
                        375
Gln Pro Glu Asn Asn Tyr Lys Thr Thr Pro Pro Val Leu Asp Ser Asp
                    390
                                        395
Gly Ser Phe Phe Leu Tyr Ser Lys Leu Thr Val Asp Lys Ser Arg Trp
                                    410
Gln Gln Gly Asn Val Phe Ser Cys Ser Val Met His Glu Ala Leu His
                                425
Asn His Tyr Thr Gln Lys Ser Leu Ser Leu Ser Pro Gly Lys
        435
                            440
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<211> 6

<212> PRT

<213> Artificial Sequence

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<223> synthetically generated peptide

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1 5

<210> 18
<211> 7
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<213> Artificial Sequence

<220>
<223> synthetically generated peptide

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1 5
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